SEQUENCE LISTING

JC20 Rec'd PCT/PTO 0 5 MAY 2009

<110> Immunokick Inc. TRANSGENIC MAMMAL CARRYING GANP GENE TRANSFERRED THEREINTO AND <120> UTILIZATION THEREOF <130> P03-0152PCT <140> PCT/JP03/014221 9 A. . --<141> 2003-11-07 <150> PCT/JP02/11598 <151> 2002-11-07 <160> 36 <170> PatentIn version 3.2 <210> 1 <211> 6429 <212> DNA <213> Mus musculus <220> <221> CDS <222> (384).. (6299) <400> 1 gttgcggtgc ggtgggcccg gtagaggctg cacgcagact gtgggcgagc acaagcgctg 60

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540	545	550

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	_		-			gca Ala						2189
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830 835	840
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	agg Arg									3245
	gtt Val					_		Arg		3293
	tgc Cys								_	3341

_	_	_												ac cca sp Pro	3389
Leu	МІА		990	Jeu 1	101	ile .		11 0. 95	נוז עו	g A.	la U	100		sp 110	
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1240

4109

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1235

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1515	1520	1525

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Phe Gly Gln Pro Ser Leu Phe Gly Gln Asn Ser Thr Pro Ser Lys Ser 35 40 45

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Val Thr Asn Ser Ser Val Thr Ser Ser Ser Phe Ile Phe Ser Lys Pro Val Thr Ser Asn Thr Pro Ala Phe Ala Ser Pro Leu Ser Asn Gln Asn Val Glu Glu Glu Lys Arg Val Ser Thr Ser Ala Phe Gly Ser Ser Asn Ser Ser Phe Ser Thr Phe Pro Thr Ala Ser Pro Gly Ser Leu Gly Glu Pro Phe Pro Ala Asn Lys Pro Ser Leu Arg Gln Gly Cys Glu Glu Ala Ile Ser Gln Val Glu Pro Leu Pro Thr Leu Met Lys Gly Leu Lys Arg Lys Glu Asp Gln Asp Arg Ser Pro Arg Arg His Cys His Glu Ala Ala Glu Asp Pro Asp Pro Leu Ser Arg Gly Asp His Pro Pro Asp Lys Arg . 305 Pro Val Arg Leu Asn Arg Pro Arg Gly Gly Thr Leu Phe Gly Arg Thr

Ile Gln Glu Val Phe Lys Ser Asn Lys Glu Ala Gly Arg Leu Gly Ser Lys Glu Ser Lys Glu Ser Gly Phe Ala Glu Pro Gly Glu Ser Asp His Ala Ala Val Pro Gly Gly Ser Gln Ser Thr Met Val Pro Ser Arg Leu Pro Ala Val Thr Lys Glu Glu Glu Glu Ser Arg Asp Glu Lys Glu Asp Ser Leu Arg Gly Lys Ser Val Arg Gln Ser Lys Arg Arg Glu Glu Trp Ile Tyr Ser Leu Gly Gly Val Ser Ser Leu Glu Leu Thr Ala Ile Gln Cys Lys Asn Ile Pro Asp Tyr Leu Asn Asp Arg Ala Ile Leu Glu Lys His Phe Ser Lys Ile Ala Lys Val Gln Arg Val Phe Thr Arg Arg Ser

Lys Lys Leu Ala Val Ile His Phe Phe Asp His Ala Ser Ala Ala Leu

Ala Arg Lys Lys Gly Lys Gly Leu His Lys Asp Val Val Ile Phe Trp His Lys Lys Lys Ile Ser Pro Ser Lys Lys Leu Phe Pro Leu Lys Glu Lys Leu Gly Glu Ser Glu Ala Ser Gln Gly Ile Glu Asp Ser Pro Phe Gln His Ser Pro Leu Ser Lys Pro Ile Val Arg Pro Ala Ala Gly Ser Leu Leu Ser Lys Ser Ser Pro Val Lys Lys Pro Ser Leu Leu Lys Met His Gln Phe Glu Ala Asp Pro Phe Asp Ser Gly Ser Glu Gly Ser Glu Gly Leu Gly Ser Cys Val Ser Ser Leu Ser Thr Leu Ile Gly Thr Val Ala Asp Thr Ser Glu Glu Lys Tyr Arg Leu Leu Asp Gln Arg Asp Arg

Ile Met Arg Gln Ala Arg Val Lys Arg Thr Asp Leu Asp Lys Ala Arg

Ala Phe Val Gly Thr Cys Pro Asp Met Cys Pro Glu Lys Glu Arg Tyr Leu Arg Glu Thr Arg Ser Gln Leu Ser Val Phe Glu Val Val Pro Gly Thr Asp Gln Val Asp His Ala Ala Ala Val Lys Glu Tyr Ser Arg Ser Ser Ala Asp Gln Glu Glu Pro Leu Pro His Glu Leu Arg Pro Ser Ala Val Leu Ser Arg Thr Met Asp Tyr Leu Val Thr Gln Ile Met Asp Gln Lys Glu Gly Ser Leu Arg Asp Trp Tyr Asp Phe Val Trp Asn Arg Thr Arg Gly Ile Arg Lys Asp Ile Thr Gln Gln His Leu Cys Asp Pro Leu Thr Val Ser Leu Ile Glu Lys Cys Thr Arg Phe His Ile His Cys Ala

His Phe Met Cys Glu Glu Pro Met Ser Ser Phe Asp Ala Lys Ile Asn

Asp Leu Arg Asn Lys Gly Val Phe Cys Ala Ser Glu Ala Glu Phe Gln Gly Tyr Asn Val Leu Leu Asn Leu Asn Lys Gly Asp Ile Leu Arg Glu Val Gln Gln Phe His Pro Asp Val Arg Asn Ser Pro Glu Val Asn Phe Ala Val Gln Ala Phe Ala Ala Leu Asn Ser Asn Asn Phe Val Arg Phe Phe Lys Leu Val Gln Ser Ala Ser Tyr Leu Asn Ala Cys Leu Leu His Cys Tyr Phe Asn Gln Ile Arg Lys Asp Ala Leu Arg Ala Leu Asn Val Ala Tyr Thr Val Ser Thr Gln Arg Ser Thr Val Phe Pro Leu Asp Gly Val Val Arg Met Leu Leu Phe Arg Asp Ser Glu Glu Ala Thr Asn Phe

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Leu Asn Tyr His Gly Leu Thr Val Ala Asp Gly Cys Val Glu Leu Asn 915 920 925

Arg Ser Ala Phe Leu Glu Pro Glu Gly Leu Cys Lys Ala Arg Lys Ser 930 935 940

Val Phe Ile Gly Arg Lys Leu Thr Val Ser Val Gly Glu Val Val Asn 945 950 955 960

Gly Gly Pro Leu Pro Pro Val Pro Arg His Thr Pro Val Cys Ser Phe.
965 970 975

Asn Ser Gln Asn Lys Tyr Val Gly Glu Ser Leu Ala Thr Glu Leu Pro 980 985 990

Ile Ser Thr Gln Arg Ala Gly Gly Asp Pro Ala Gly Gly Gly Arg Gly 995 1000 1005

Glu Asp Cys Glu Ala Glu Val Asp Leu Pro Thr Leu Ala Val Leu 1010 1015 1020

Pro Gln Pro Pro Pro Ala Ser Ser Ala Thr Pro Ala Leu His Val 1025 1030 . 1035

Gln Pro Leu Ala Pro Ala Ala Pro Ser Leu Leu Gln Ala Ser 1040 1045 1050

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- Val Ala Ala Ala Leu Gly Val Ser Asn Ala Ala Val Glu Asp Leu 1100 1105 1110
- Ile Thr Ala Ala Thr Thr Gly Ile Leu Arg His Val Ala Ala Glu 1115 1120 1125
- Glu Val Ser Met Glu Arg Gln Arg Leu Glu Glu Glu Lys Gln Arg 1130 1135 1140
- Ala Glu Glu Glu Arg Leu Lys Gln Glu Arg Glu Leu Met Leu Thr 1145 1150 1155
- Gln Leu Ser Glu Gly Leu Ala Ala Glu Leu Thr Glu Leu Thr Val 1160 1165 1170
- Thr Glu Cys Val Trp Glu Thr Cys Ser Gln Glu Leu Gln Ser Ala 1175 1180 1185

- Val Lys Ile Asp Gln Lys Val Arg Val Ala Arg Cys Cys Glu Ala 1190 1195 1200
- Val Cys Ala His Leu Val Asp Leu Phe Leu Ala Glu Glu Ile Phe 1205 1210 1215
- Gln Thr Ala Lys Glu Thr Leu Gln Glu Leu Gln Cys Phe Cys Lys 1220 1225 1230
- Tyr Leu Gln Arg Trp Arg Glu Ala Val Ala Ala Arg Lys Lys Phe 1235 1240 1245
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- Asn Lys Thr Ala His Gln Ile Lys Val Gln His Phe His Gln Gln 1310 1315 1320

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- Thr Val Ser Val Asn Val Cys Ile Lys Val Ala His Gly Thr Leu 1415 1420 1425
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- Gly Thr Ser Gly Leu Met Leu Leu Leu Pro Pro Lys Val Lys Ser 1445 1450 1455

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- Glu Glu Val Ala Glu Glu Glu Leu Ser Trp Leu Ser Ala Leu Leu 1460 1465 1470
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- Lys Leu Ile Ser Asp Tyr Ile Val Val Glu Ile Pro Asp Ser Val 1520 1525 1530
- Asn Asp Leu Gln Gly Thr Val Lys Val Ser Gly Ala Val Gln Trp 1535 1540 1545
- Leu Ile Ser Gly Cys Pro Gln Ala Leu Asp Leu Cys Cys Gln Thr 1550 1555 1560
- Leu Val Gln Tyr Val Glu Asp Gly Ile Ser Arg Glu Phe Ser Arg 1565 1570 1575
- Arg Phe Phe His Asp Arg Arg Glu Arg Arg Leu Ala Ser Leu Pro 1580 1585 1590

- Ser Gln Glu Pro Ser Thr Ile Ile Glu Leu Phe Asn Ser Val Leu 1595 1600 1605
- Gln Phe Leu Ala Ser Val Val Ser Ser Glu Gln Leu Cys Asp Ile 1610 1615 1620
- Ser Trp Pro Val Met Glu Phe Ala Glu Val Gly Gly Ser Gln Leu 1625 1630 1635
- Leu Pro His Leu His Trp Asn Ser Pro Glu His Leu Ala Trp Leu 1640 1645 1650
- Lys Gln Ala Val Leu Gly Phe Gln Leu Pro Gln Met Asp Leu Pro 1655 1660 1665
- Pro Pro Gly Ala Pro Trp Leu Pro Val Cys Ser Met Val Ile Gln 1670 1675 1680
- Tyr Thr Ser Gln Ile Pro Ser Ser Ser Gln Thr Gln Pro Val Leu 1685 1690 1695
- Gln Ser Gln Ala Glu Asn Leu Leu Cys Arg Thr Tyr Gln Lys Trp 1700 1705 1710
- Lys Asn Lys Ser Leu Ser Pro Gly Gln Glu Leu Gly Pro Ser Val 1715 1720 1725

- Ala Glu Ile Pro Trp Asp Asp Ile Ile Thr Leu Cys Ile Asn His 1730 1735 1740
- Lys Leu Arg Asp Trp Thr Pro Pro Arg Leu Pro Val Thr Leu Glu 1745 1750 1755
- Ala Leu Ser Glu Asp Gly Gln Ile Cys Val Tyr Phe Phe Lys Asn 1760 1765 1770
- Leu Leu Arg Lys Tyr His Val Pro Ser Ser Trp Glu Gln Ala Arg 1775 1780 1785
- Met Gln Thr Gln Arg Glu Leu Gln Leu Ser His Gly Arg Ser Gly 1790 1795 1800
- Met Arg Ser Ile His Pro Pro Thr Ser Thr Phe Pro Thr Pro Leu 1805 1810 1815
- Leu His Val His Gln Lys Gly Lys Lys Glu Glu Ser Gly Arg 1820 1825 1830
- Glu Gly Ser Leu Ser Thr Glu Asp Leu Leu Arg Gly Ala Ser Ala 1835 1840 1845
- Glu Glu Leu Leu Ala Gln Ser Leu Ser Ser Ser Leu Leu Glu Glu 1850 1855 1860

Lys Glu Glu Asn Lys Arg Phe Glu Asp Gln Leu Gln Gln Trp Leu 1865 1870 1875

Ser Gln Asp Ser Gln Ala Phe Thr Glu Ser Thr Arg Leu Pro Leu 1880 1885 1890

Tyr Leu Pro Gln Thr Leu Val Ser Phe Pro Asp Ser Ile Lys Thr 1895 1900 1905

Gln Thr Met Val Lys Thr Ser Thr Ser Pro Gln Asn Ser Gly Thr 1910 1915 1920

Gly Lys Gln Leu Arg Phe Ser Glu Ala Ser Gly Ser Ser Leu Thr 1925 1930 1935

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Glu Glu Ala Ala Ser Glu Leu His Leu Ser Ala Leu Leu Glu Met 1955 1960 1965

Val Asp Met 1970

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ttc	agt	ggg	cag	cag	cct	agt	gct	ttt	tcg	gcg	tct	tct	agt	aat	gta	103
Phe	Ser	Gly	Gln	Gln	Pro	Ser	Ala	Phe	Ser	Ala	Ser	Ser	Ser	Asn	Val	
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Gly	Thr	Leu	Pro	Ser	Lys	Pro	Pro	Phe	Arg	Phe	Gly	Gln	Pro	Ser	Leu	
		25					30					35				
																100
								ggg								199
Phe		Gln	Asn	Ser	Thr		Ser	Gly	Lys	Ser		Gly	Phe	Ser	GIn	
	40					45					50					
									4			+	+ 0 +	+ 0.0	at a	247
								gga								241
	Ser	Ser	Phe	Pro		Ser	Ser	Gly	vai	65	птѕ	261	Ser	261	70	
55					60					00					10	
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ctt	gag	cac	act	too	acc	ttt	gtg	gct	acc	tct	ggg	cct	tca	agt	tca	343
															Ser	
200			90					95			·		100			

gtg Val									391
gct Ala 120									439
aac Asn									487
gca Ala									535
cag Gln	_		-						583
agt Ser				-					631
agt Ser 200									679
aat Asn									727
gag Glu									775

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								gaa Glu		871
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_								cca Pro		967
								aaa Lys		1015
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_	_	_						ggc Gly		1111
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		ggc Gly							1447
		ctt Leu							1495
		aag Lys							1543
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			acc Thr]	1975
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			gac Asp							2	2071

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Ser	Ser	Ala	Asp	Gln	Glu	Glu	Pro	Leu	Pro	His	Glu	Leu	Arg	Pro	Leu	
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Pro	Val	Leu	Ser	Arg	Thr	Met	Asp	Tyr	Leu		Thr	Gln	Ile	Met		
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Gln	Lys	Glu	Gly		Leu	Arg	Asp	Trp		Asp	Phe	Val	irp	725	Arg	
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					Lys											
	,,, B	01)	730	6	2,0	,		735					740	•		
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ctg	acg	gtg	tcc	ctg	att	gag	aag	tgc	acc	cgg	ttt	cac	atc	cac	tgt	2311
Leu	Thr	Val	Ser	Leu	Ile	Glu	Lys	Cys	Thr	Arg	Phe	His	Ile	His	Cys	
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					gag											2359
Ala		Phe	Met	Cys	Glu		Pro	Met	Ser	Ser		Asp	Ala	Lys	Ile	
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	. 4			_4_			4	a+ a	20.5	0.70	0 t a	000	ana	a t a	tac	2407
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775	ASII	GIU	ASII	мес	780	Lys	Cys	Leu	0111	785	Leu	Lys	Olu	Mec	790	
773					100					, 00	٠				, , ,	
cag	gac	ctg	aga	aac	aag	ggt	gtc	ttc	tgt	gcc	agc	gaa	gcg	gag	ttc	2455
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Gln	Gly	Tyr	Asn	Val	Leu	Leu	Ser	Leu	Asn	Lys	Gly	Asp	Ile	Leu	Arg	
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		ttt Phe						2599
		cag Gln 860						2647
		cag Gln						2695
		agc Ser						2743
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		ggc Gly						2839
		ctg Leu 940						2887
		agg Arg						2935

		Gly I		cc gt Pro Va		o Ar			Cys	2983
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	-	Ser		ccc Pro 1005						3076
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	cca Pro 1030	G1n		gcc Ala 1035						3166
		Val		acc Thr 1050				_		3211
	ctg Leu 1060	Ser		gaa Glu 1065					gtg Val	3256
		Ser		ctg Leu 1080						3301
	gag Glu 1090	Ala		gac Asp 1095						3346

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Glu	Asp	Leu	Leu	Thr	Ala		Thr	Thr	Gly	He		Arg	His	He	
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Val	Leu	Ser	Glu	Leu	Ser	Gln	Gly	Leu	Ala	Val	Glu	Leu	Met	Glu	
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Arg	Val	Met	Met	Glu	Phe	Val	Arg	Glu	Thr	Cys			Glu	Leu	
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											4		. ~ +	+ ~ ~	3661
	aat														3001
Lys	Asn	Ala	vaı	Glu	Inr			Arg	vai	Arg	1205	нта	MI B	Cys	
	1195					1200					1203				
t o t	gag	σat	gtc	tøt.	gcc	cac	tta	gtg	gac	ttg	ttt	ctc	gtg	gag	3706
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Phe	Cys	Lys	Tyr	Leu	Gln	Arg	Trp	Arg	Glu	Ala	Val	Thr	Ala	Arg	
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Glu	Cys	Pro	Ile	Ala	Glu	Glu	Asn	Leu	Ala	Arg	Gly	Leu	Leu	Asp	
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Leu	Gly	His	Ala	Gly	Arg	Leu	Gly	Ile	Ser	Cys		Arg	Leu	Arg	
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Arg		Arg	Asn	Lys	ınr	Ala 1320	HIS	GIII	меι	Lys	1325		1112	The	•
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Tyr	Gln	G1n	Leu	Leu	Ser	Asp	Val	Ala	Trp	Ala	Ser	Leu	Asp	Leu	
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Gly	Ala	Leu	Ser	Asp	Gly	Ala	Ile	Asp	Ala	Val	Glu	Thr	Gln	Lys	
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Met	Lys	Ser	Glu	Asp	Met	Ala	Glu	Glu	Asp	Val			Leu	Ser	
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		_		cct Pro 1605					4876
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		Glu			gaa Glu 1770	Asp			Val		5371

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				cag Gln 1800					5461
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		_		cac His 1830					5551
				att Ile 1845					5596
				ctc Leu 1860					5641
				gag Glu 1875					5686
cag Gln				gac Asp 1890					5731
				cct Pro 1905					5776

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Thr Il	e Glu	Pro	Val	Met	Lys	Thr	Ser	Val	Thr	Thr	Ser	Pro	Gln	
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		-												
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Ser As	p Met	Met	Arg	Glu	Gln	Leu	Gln	Leu	Ser	Glu	Ala	Thr	Gly	
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acg tg														5911
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agt tc					gtt									5950
Ser Se	_	GIU	GIU	GIU	Val 1965	нта	Ser	Glu	Leu	1970	Leu	361	nia	
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Leu Le		Met				-0-0	36		,	-0-00	5 0-1	5000		
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Ser Phe Lys Pro Leu Glu Asn Ala Val Phe Lys Pro Ile Leu Gly Ala

Glu Ser Glu Pro Glu Lys Thr Gln Ser Gln Ile Ala Ser Gly Phe Phe 165 170 175

Thr Phe Ser His Pro Ile Ser Ser Ala Pro Gly Gly Leu Ala Pro Phe 180 185 190

Ser Phe Pro Gln Val Thr Ser Ser Ser Ala Thr Thr Ser Asn Phe Thr 195 200 205

Phe Ser Lys Pro Val Ser Ser Asn Asn Ser Leu Ser Ala Phe Thr Pro 210 215 220

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Ile Phe Gly Ser Ser Asn Asn Ser Phe Ser Ser Phe Pro Val Ser Ser 245 250 255

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Gly Cys Glu Glu Ala Val Ser Gln Val Glu Pro Leu Pro Ser Leu Met 275 280 285

Lys Gly Leu Lys Arg Lys Glu Asp Gln Asp Arg Ser Pro Arg Arg His 290 295 300 Gly His Glu Pro Ala Glu Asp Ser Asp Pro Leu Ser Arg Gly Asp His 305 310 315 320

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Gly Arg Leu Gly Asn Lys Glu Ala Lys Lys Glu Thr Gly Phe Val Glu 355 360 365

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- Phe Thr Asp Leu Thr Ser Leu Pro Leu Tyr Leu Pro Gln Thr Leu 1895 1900 1905
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